

WE CLAIM:

1. A pharmaceutical preparation for administration to a female in need of hormone replacement therapy, comprising a plurality of doses for administration in alternating relatively dominant estrogenic activity phases, the relatively dominant estrogenic activity phases consisting of a relatively dominant estrogenic activity relatively dominant estrogenic activity phase comprising three consecutive daily doses or an equivalent thereof, of a substance exhibiting estrogenic activity equivalent to about 1 mg per day of 17 β -estradiol, and a relatively dominant progestogenic activity phase comprising three consecutive daily doses or an equivalent thereof, of a combination of a substance exhibiting estrogenic activity equivalent to about 1 mg per day of 17 β -estradiol and a substance exhibiting progestogenic activity equivalent to about 90 μ g per day of norgestimate.

2. The pharmaceutical preparation as claimed in claim 1 for administration to a female in need of hormone replacement therapy comprising a plurality of daily doses for consecutive administration, the doses being administered in alternating phases, the phases comprising a relatively dominant estrogenic activity phase comprising three consecutive daily doses of a substance exhibiting estrogenic activity equivalent to about 1 mg per day of 17 β -estradiol, and a relatively dominant progestogenic activity phase comprising three consecutive daily doses of a combination of a substance. exhibiting estrogenic activity equivalent to about 1 mg per day of 17 β -estradiol and a substance exhibiting progestogenic activity equivalent to about 90 μ g per day of norgestimate.

3. The pharmaceutical preparation as claimed in claim 2 wherein the substance exhibiting progestogenic activity is selected on the basis that it binds to progestin receptors, demonstrates poor affinity for androgen receptors and has a lack of affinity for sex-hormone-binding globulin.

4. The pharmaceutical preparation as claimed in claim 3 wherein the substance exhibiting estrogenic activity is selected from 17 α -ethinylestradiol, esters and ethers of 17 α -ethinylestradiol, 17 α -ethinylestradiol 3-dimethylamino propionate, 17 α ethinylestradiol 3-cyclopentyl ether (quienestrol) and 17 α -ethinylestradiol 3-methyl ether (mestranol); natural estrogens, estrone, estrone sulfate, estrone sulfate piperazine salt, estradiol and estriol, and their esters, as well as the synthetic estrogens, and the substance exhibiting progestogenic activity

is selected from desogestrel, dydrogesterone, medroxyprogesterone acetate, norethynodrel, cyproterone acetate, chlormadinone acetate, magestrol acetate, 17 D-acetyl norgestimate, dienogest, trimegestone, drospirinone and nomagestrel.

5 5. The pharmaceutical preparation as claimed in claim 1 wherein the doses are in oral form.

6. The pharmaceutical preparation as claimed in claim 5 wherein the oral form is a tablet.

10 7. A package containing a pharmaceutical preparation for administration to a female in need of hormone replacement therapy, comprising a plurality of doses arranged in alternating phases, the phases consisting of a relatively dominant estrogenic activity phase comprising three consecutive daily doses or an equivalent
15 thereof, of a substance exhibiting estrogenic activity equivalent to about 1 mg per day of 17 β -estradiol, and a relatively dominant progestogenic activity phase comprising three consecutive daily` doses or an equivalent thereof, of a combination of a substance exhibiting estrogenic activity equivalent to about 1 mg per day of 17 β -estradiol and a. substance exhibiting progestogenic activity
20 equivalent to about 90 μ g per day of norgestimate.

8. The pharmaceutical package as claimed in claim 7 containing a pharmaceutical regimen for administration to a female in need of hormone replacement therapy, the doses being arranged for administration in alternating
25 phases, the phases consisting of a relatively dominant estrogenic activity phase comprising three consecutive daily doses of a substance exhibiting estrogenic activity equivalent to about 1 mg per day of 17 β -estradiol, and a relatively dominant progestogenic activity phase comprising three consecutive daily doses comprising a combination of a substance exhibiting estrogenic activity equivalent to
30 about 1 mg per day of 17 β -estradiol and a substance exhibiting progestogenic activity equivalent to about 90 μ g per day of norgestimate.

9. The pharmaceutical package as claimed in claim 8 wherein the substance exhibiting progestogenic activity is selected on the basis that it binds to progesterin
35 receptors, it demonstrates poor affinity for androgen receptors and has a lack of affinity for sex-hormone-binding globulin.

10. The pharmaceutical package as claimed in claim 8 wherein the substance exhibiting estrogen activity is selected from 17 α -ethinylestradiol, esters and ethers of 17 α -ethinylestradiol, 17 α -ethinylestradiol 3-dimethylamino propionate, 17 α -ethinylestradiol 3-cyclopentyl ether (quienestrol) and. 17 α -ethinylestradiol 3-methyl
5 ether (mestranol); natural estrogens, estrone, estrone sulfate, estrone sulfate piperazine salt, estradiol and estriol, and their esters, as well as the synthetic estrogens, and the substance exhibiting progestogenic activity is selected from desogestrel, dydrogesterone, medroxyprogesterone acetate, norethynodrel, cyproterone acetate, chlormadinone acetate, magestrol acetate, 17 D-acetyl
10 norgestimate, dienogest, trimegestone, drospirinone and nomagestrel.

11. The pharmaceutical package as claimed in claim 7, wherein the doses are in oral form.

12. The pharmaceutical package as claimed in claim 11, wherein the oral
15 form is a tablet.

13. A method of treating a female in need of hormone replacement therapy comprising administering to said female a pharmaceutical regimen comprising a
20 plurality of doses arranged in alternating phases, the phases comprising a relatively dominant estrogenic activity phase comprising three consecutive daily doses or an equivalent thereof, of a substance exhibiting estrogenic activity equivalent to about 1 mg per day of 17 β -estradiol, and a relatively dominant progestogenic activity phase comprising three consecutive daily doses or an
25 equivalent thereof, of a combination of a substance exhibiting estrogenic activity equivalent to about 1 mg per day of 17 β -estradiol and a substance exhibiting progestogenic activity equivalent to about 90 μ g per day of norgestimate.

14. A method of treating a female in need of hormone replacement therapy as
30 claimed in claim 13 comprising administering to said female a pharmaceutical regimen comprising a plurality of doses arranged in alternating phases, the phases comprising a relatively dominant estrogenic activity phase comprising three daily doses of a substance exhibiting estrogenic activity equivalent to about 1 mg per day of 17 β -estradiol, and a relatively dominant progestogenic activity phase comprising
35 three daily doses of a combination of a substance exhibiting estrogenic activity equivalent to about 1 mg per day of 17 β -estradiol and a substance exhibiting

progestogenic activity equivalent to about 90 µg per day of norgestimate.

15. The method as claimed in claim 14 wherein the substance exhibiting progestogenic activity is selected on the basis that it binds to progestogen
5 receptors, demonstrates poor affinity for androgen receptors and has a lack of affinity for sex-hormone-binding globulin.

16. The method as claimed in claim 14 wherein the substance exhibiting estrogenic activity is selected from 17α-ethinylestradiol, esters and ethers of 17α-
10 ethinylestradiol, 17α-ethinylestradiol 3-dimethyl amino propionate, 17α-ethinylestradiol 3-cyclopentyl ether (quienestrol) and 17α-ethinylestradiol 3-methyl ether (mestranol); natural estrogens, estrone, estrone sulfate, estrone sulfate piperazine salt, estradiol and estriol, and their esters, as well as the synthetic estrogens, and the substance exhibiting progestogenic activity is selected from
15 desogestrel, dydrogesterone, medroxyprogesterone acetate, norethynodrel, cyproterone acetate, chlormadinone acetate, magesrol acetate, 17 D-acetyl norgestimate, dienogest, trimegestone, drospirinone and nomagestrel.

17. The method as claimed in claim 13 wherein the doses are in oral form.

18. The method as claimed in claim 17 wherein the oral form is a tablet.

19. A use of estrogenically active substance and a progestogenically active substance in the preparation of a medicament, characterized in that the
25 medicament is for hormone replacement therapy for administration to a female in need of such therapy, the medicament comprising a plurality of doses for consecutive administration in alternating phases, which phases consist of a relatively dominant estrogenic activity phase comprising three consecutive daily doses or an equivalent thereof, of a substance exhibiting estrogenic activity
30 equivalent to about 1 mg per day of 17β-estradiol, and a relatively dominant progestogenic activity phase comprising three daily doses or an equivalent thereof, of a combination of a substance exhibiting estrogenic activity equivalent to about 1 mg per day of 17β-estradiol and a substance exhibiting progestogenic activity equivalent to about 90 µg per day of norgestimate.

20. The use as claimed in claim 19 wherein the medicament comprises a

plurality of doses for consecutive administration in alternating phases, which phases consist of a relatively dominant estrogenic activity phase comprising three consecutive daily doses of a substance exhibiting estrogenic activity equivalent to about 1 mg per day of 17 β -estradiol, and a relatively dominant progestogenic activity phase comprising three daily doses of a combination of a substance exhibiting estrogenic activity equivalent to about 1 mg per day of 17 β -estradiol and a substance exhibiting progestogenic activity equivalent to about 90 μ g per day of norgestimate.

21. The use as claimed in claim 20 wherein the substance exhibiting progestogenic activity is selected on the basis that it binds to progestin receptors, it demonstrates poor affinity for androgen receptors and has a lack of affinity for sex-hormone-binding globulin.

22. The use as claimed in claim 20 wherein the substance exhibiting estrogenic activity is selected from 17 α -ethinylestradiol 3-methyl ether (mestranol); natural estrogens, estrone, estrone sulfate, estrone sulfate piperazine salt, estradiol and estriol, and their esters, as well as the synthetic estrogens, and the substance exhibiting progestogenic activity is selected from desogestrel, dydrogesterone, medroxyprogesterone acetate, norethynodrel, cyproterone acetate, chlormadinone acetate, magestrol acetate, 17D-acteyl norgestimate, dienogest, trimegestone, dropserinone and nomagestrel.

23. The preparation as claimed in claim 1 in which the daily doses in the relatively dominant estrogenic activity phase comprise about 1mg per day. of 17 β -estradiol, and in the relatively dominant progestogenic activity phase comprise about 1mg per day of 17 β -estradiol and about 90 μ g per day of norgestimate.

24. The preparation as claimed in claim 23 wherein the doses are in oral form.

25. The preparation as claimed in claim 24 wherein the doses are in tablet form.

26. The package claimed in claim 7 in which the daily doses in the relatively

dominant estrogenic activity phase comprise about 1mg per day of 17 β -estradiol and in the relatively dominant progestogenic activity phase comprise about 1 mg per day of 17 β -estradiol and about 90 μ g per day of norgestimate.

27. The package as claimed in claim 26 wherein the doses are in oral form.

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28. The package as claimed in claim 27 wherein the doses are in tablet form.

29. The method as claimed in claim 14 in which the daily doses in the relatively dominant estrogenic activity phase comprise about 1 mg per day of 17 β -estradiol, and in the relatively dominant progestogenic activity phase comprise about 1 mg per day of 17 β -estradiol and about 90 μ g per day of norgestimate.

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30. The method as claimed in claim 29 wherein the doses are in oral form.

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31. The method as claimed in claim 30 wherein the doses are in tablet form.

32. The use as claimed in claim 20 in which the daily doses in the relatively dominant estrogenic activity phase comprises about 1 mg per day of 17 β -estradiol, and in the relatively dominant progestogenic activity phase comprises about 1 mg per day of 17 β -estradiol and about 90 μ g per day of norgestimate.

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33. The use as claimed in claim 32 wherein the doses are in oral form.

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34. The use as claimed in claim 33 wherein the doses are in tablet form.